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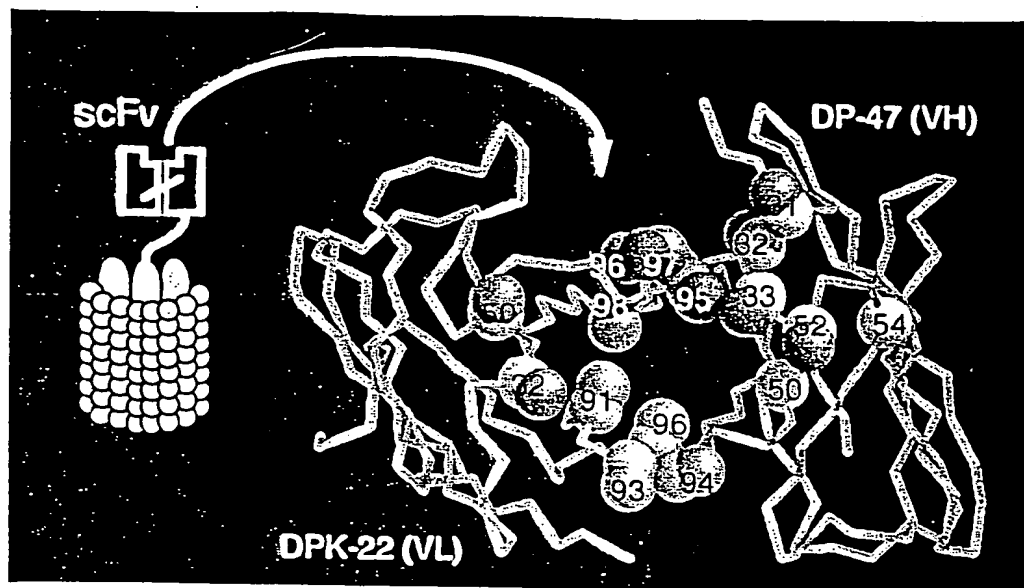
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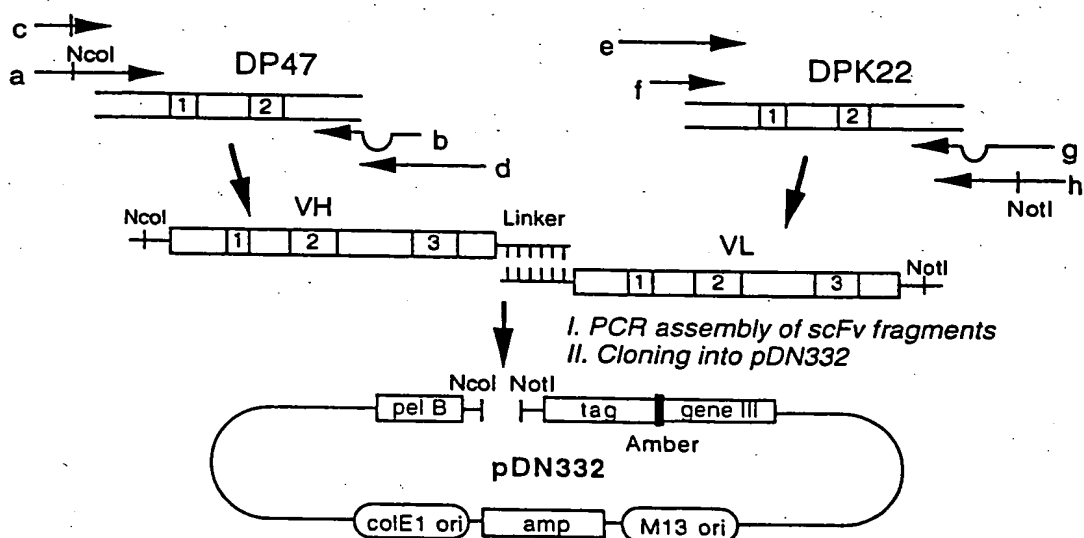
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A)



B)

VH primers:

| | | |
|---|-----------|--|
| a | DP47baNco | GCG GCC CAG CAT GCC ATG GCC GAG GTG CAG CTG TTG GAG TCT GGG |
| b | CDR3for | GGT TCC CTG GCC CCA GTA GTC AAA MNN MNN MNN MNN TTT CGC ACA GTA ATA TAC G |
| c | VHpulth | GCG GCC CAG CAT GCC ATG GCC GAG |
| d | Jassm | CCC GCT ACC GCC ACT GGA CCC ATC GCC ACT CGA GAC GGT GAC CAG GGT TCC CTG GCC CCA GTA GTC |

VL primers:

| | | |
|---|-----------|--|
| e | DPK22assm | GAT GGG TCC AGT GGC GGT AGC GGG GGC GCG TCG ACT GGC GAA ATT GTG TTG ACG CAG TCT CC |
| f | DPK3for | CAC CTT GGT CCC TTG GCC GAA CGT MNN CGG MNN MNN ACC MNN CTG CTG ACA GTA ATA CAC TGC |
| g | Jfornot | GAG TCA TTC TCG ACT TGC GGC CGC TTT GAT TTC CAC CTT GGT CCC TTG GCC GAA CG |
| h | pulth | GAT GGG TCC AGT GGC GGT AGC GGG |

Figure 1

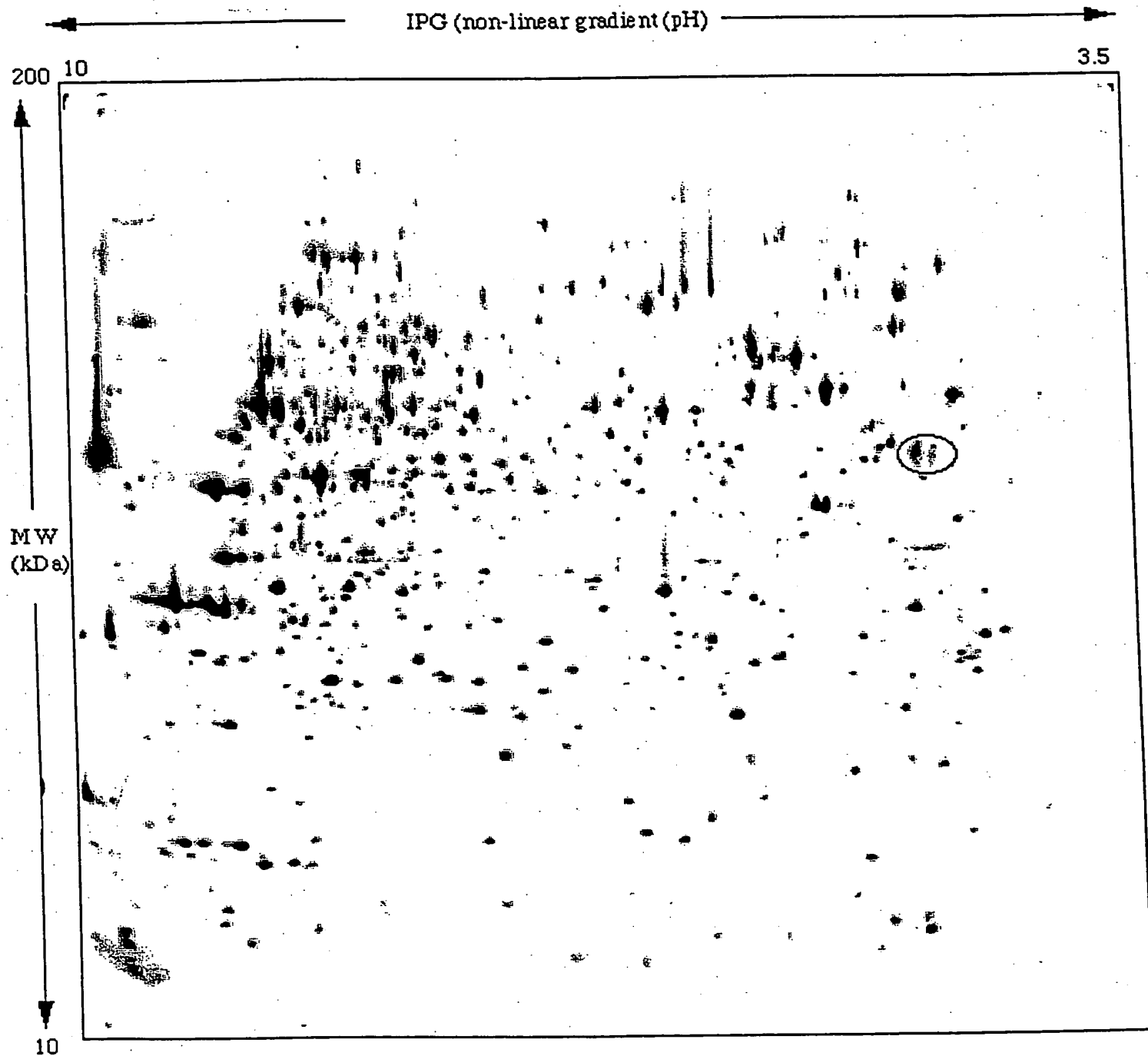


Figure 2A

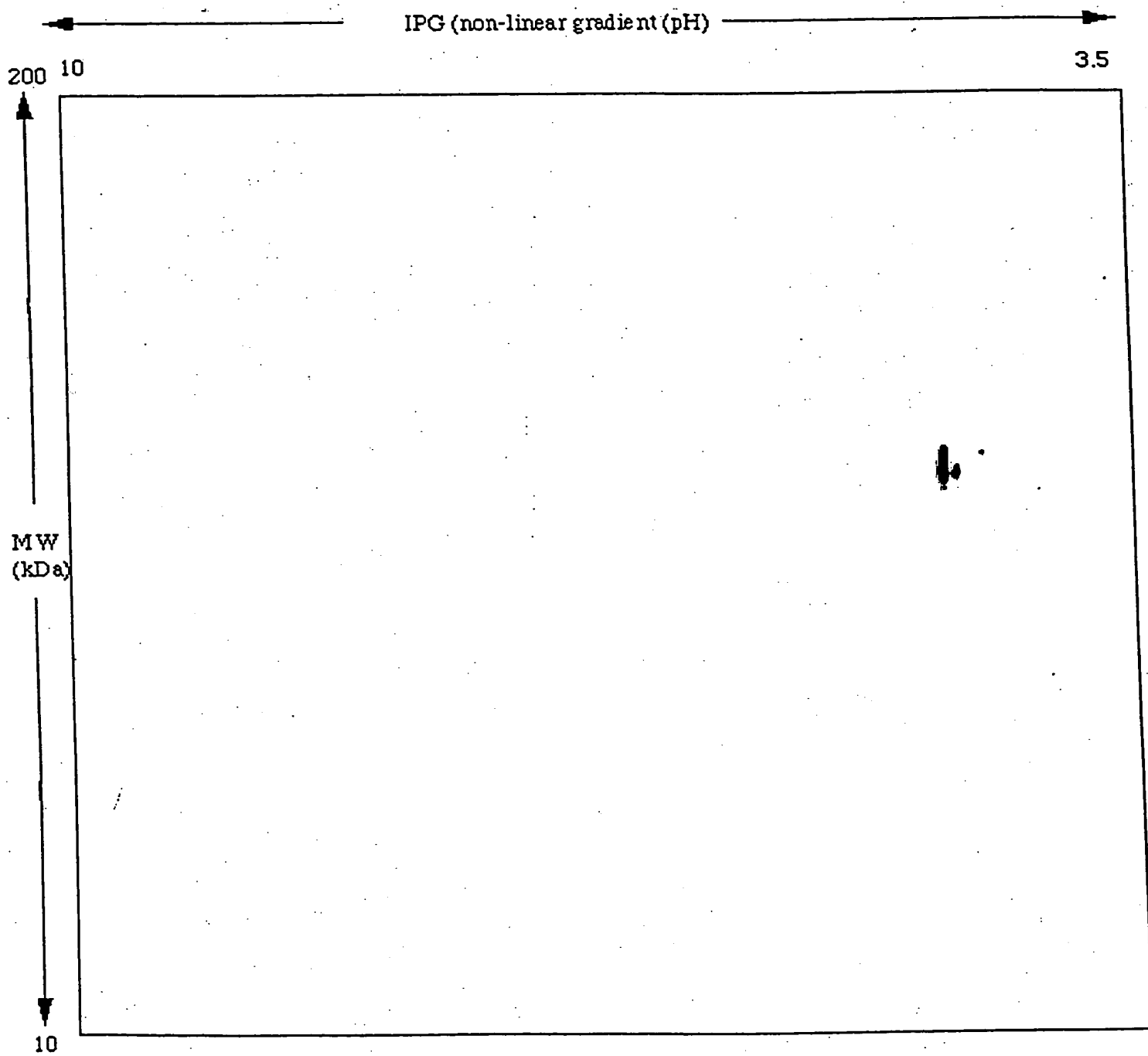


Figure 2B



Figure 3

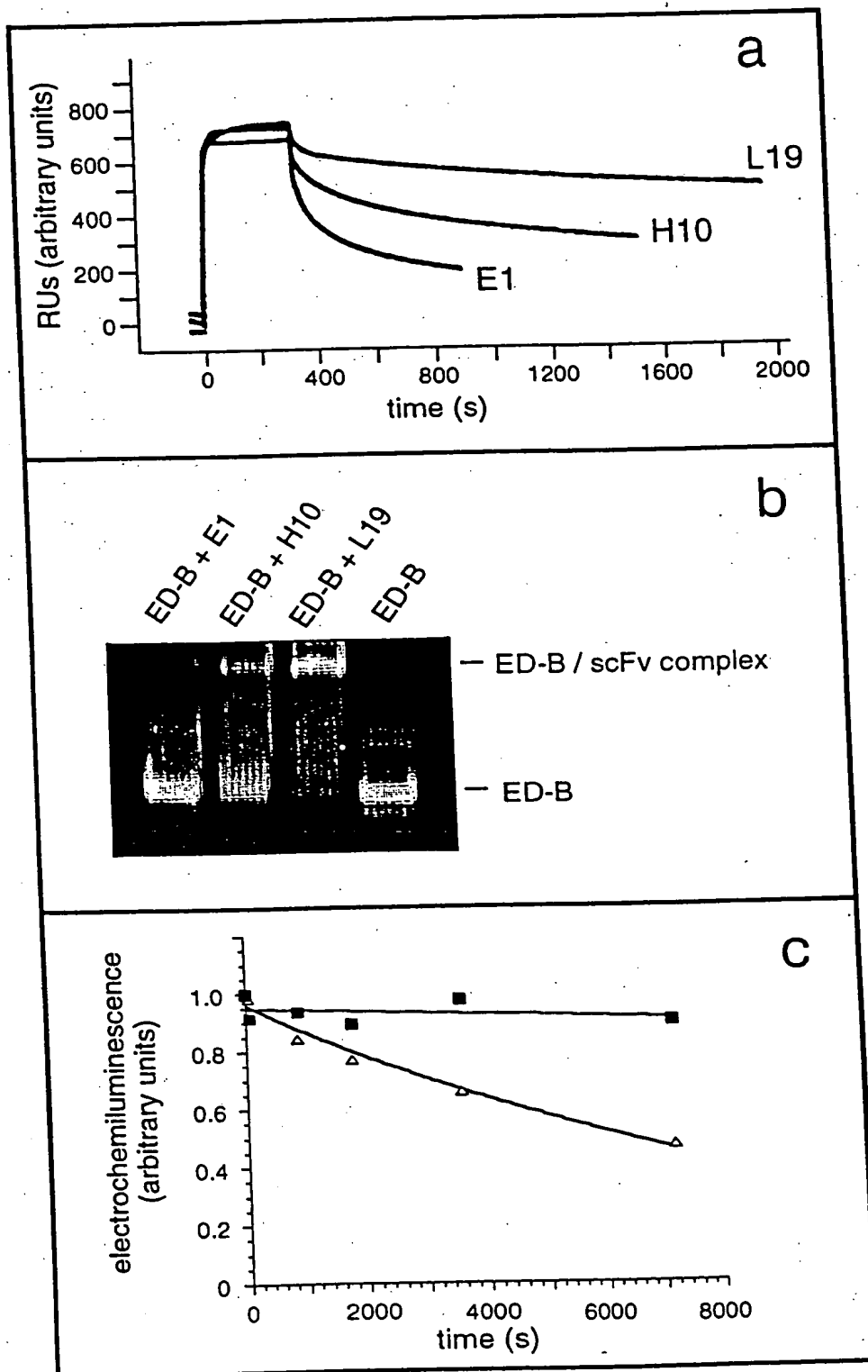
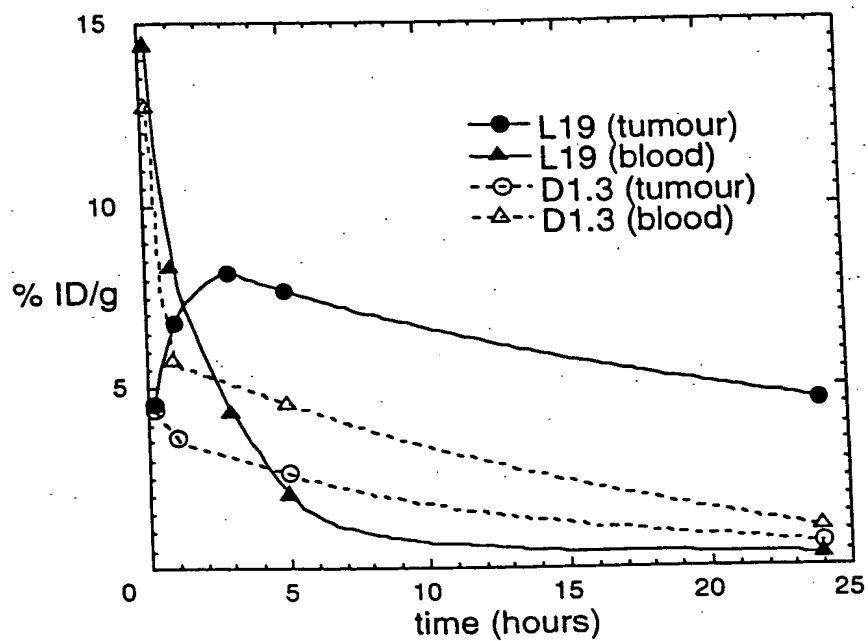


Figure 4



| time (h) \ %ID/g | scFv (L19) | | | | | | | scFv (D1.3) | |
|------------------|------------|-----------|------------|------------|-------------|------------|-----------|-------------|-----------|
| | Kidney | Spleen | Lung | Liver | Brain | Blood | Tumor | Blood | Tumor |
| 0.25 | 26.4 ± 4.7 | 5.4 ± 0.4 | 15.1 ± 5.1 | 5.7 ± 0.6 | 0.4 ± 0.04 | 14.4 ± 2.6 | 4.6 ± 1.0 | 12.7 ± 0.1 | 4.4 ± 0.4 |
| 1 | 19.2 ± 3.9 | 3.8 ± 0.3 | 9.8 ± 0.9 | 2.8 ± 0.3 | 0.3 ± 0.1 | 8.3 ± 0.9 | 6.9 ± 2.4 | 5.7 ± 0.7 | 3.6 ± 1.0 |
| 3 | 8.1 ± 1.6 | 2.0 ± 0.3 | 5.0 ± 1.4 | 1.7 ± 0.02 | 0.2 ± 0.01 | 4.3 ± 0.3 | 8.2 ± 4.2 | — | — |
| 5 | 4.2 ± 1.1 | 1.8 ± 0.2 | 3.5 ± 0.2 | 1.3 ± 0.3 | 0.1 ± 0.02 | 2.0 ± 1.6 | 7.7 ± 2.5 | 4.6 ± 1.2 | 2.6 ± 1.5 |
| 24 | 0.7 ± 0.1 | 0.4 ± 0.1 | 1.0 ± 0.1 | 0.2 ± 0.04 | 0.02 ± 0.01 | 0.4 ± 0.1 | 4.7 ± 0.6 | 1.1 ± 0.5 | 0.7 ± 0.4 |

Figure 5

VH

| | | |
|---------------------|---------------------|---------------------|
| E V Q L L E S G G G | L V Q P G G S L R L | S C A A S G F T F S |
| S F S M S W V R Q A | P G K G L E W V S S | I S G S S G T T Y Y |
| A D S V K G R F T I | S R D N S K N T L Y | L Q M N S L R A E D |
| T A V Y Y C A K P F | P Y F D Y W G Q G T | L V T V S S |

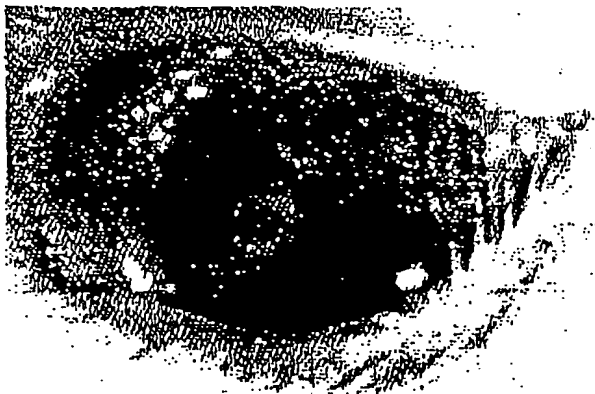
linker

G D G S S G G S G G A S T G

VL

| | | |
|---------------------|---------------------|---------------------|
| E I V L T Q S P G T | L S L S P G E R A T | L S C R A S Q S V S |
| S S Y L A W Y Q Q K | P G Q A P R L L I Y | Y A S S R A T G I P |
| D R F S G S G S G T | D F T L T I S R L E | P E D F A V Y Y C Q |
| Q T G R I P P T F G | Q G T K V E I K | |

Figure 6: Amino acid sequence of antibody L19



day 9 / rabbit 1 / left / nC



day 9 / rabbit 1 / right / VEGF high



day 9 / rabbit 2 / left / PMA



day 9 / rabbit 2 / left / VEGF low



day 9 / rabbit 3 / left / VEGF high



day 9 / rabbit 3 / right / PMA



day 9 / rabbit 4 / left / PMA



day 9 / rabbit 4 / right / nC

Figure 7

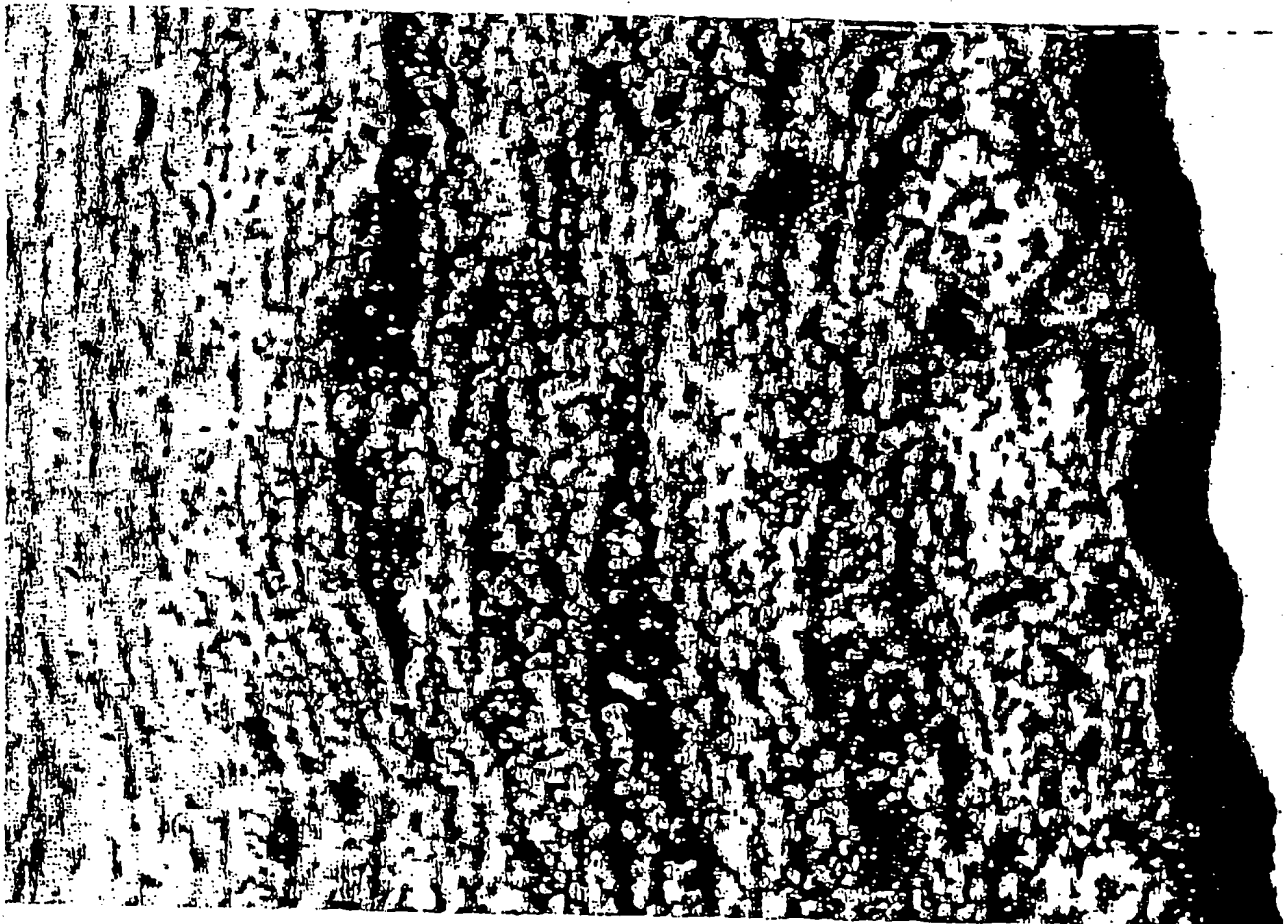


Figure 8

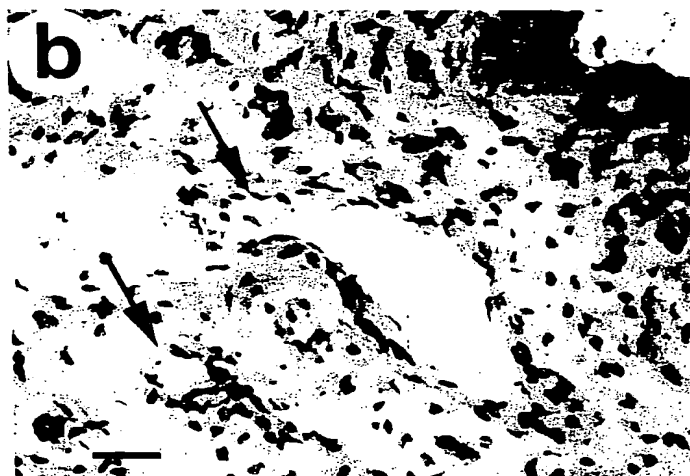
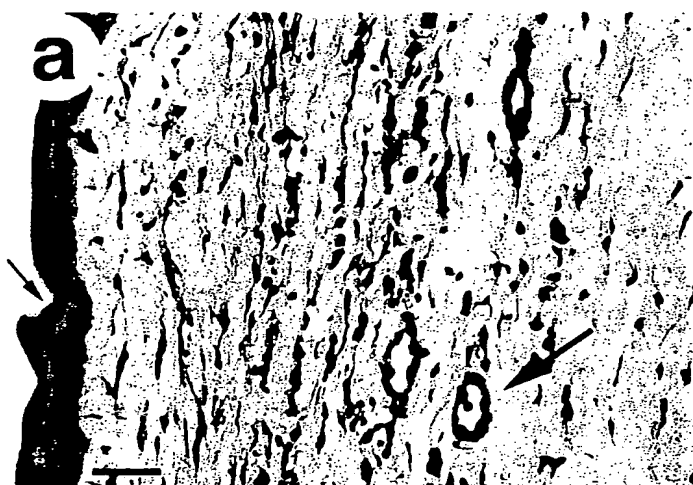


FIG. 9

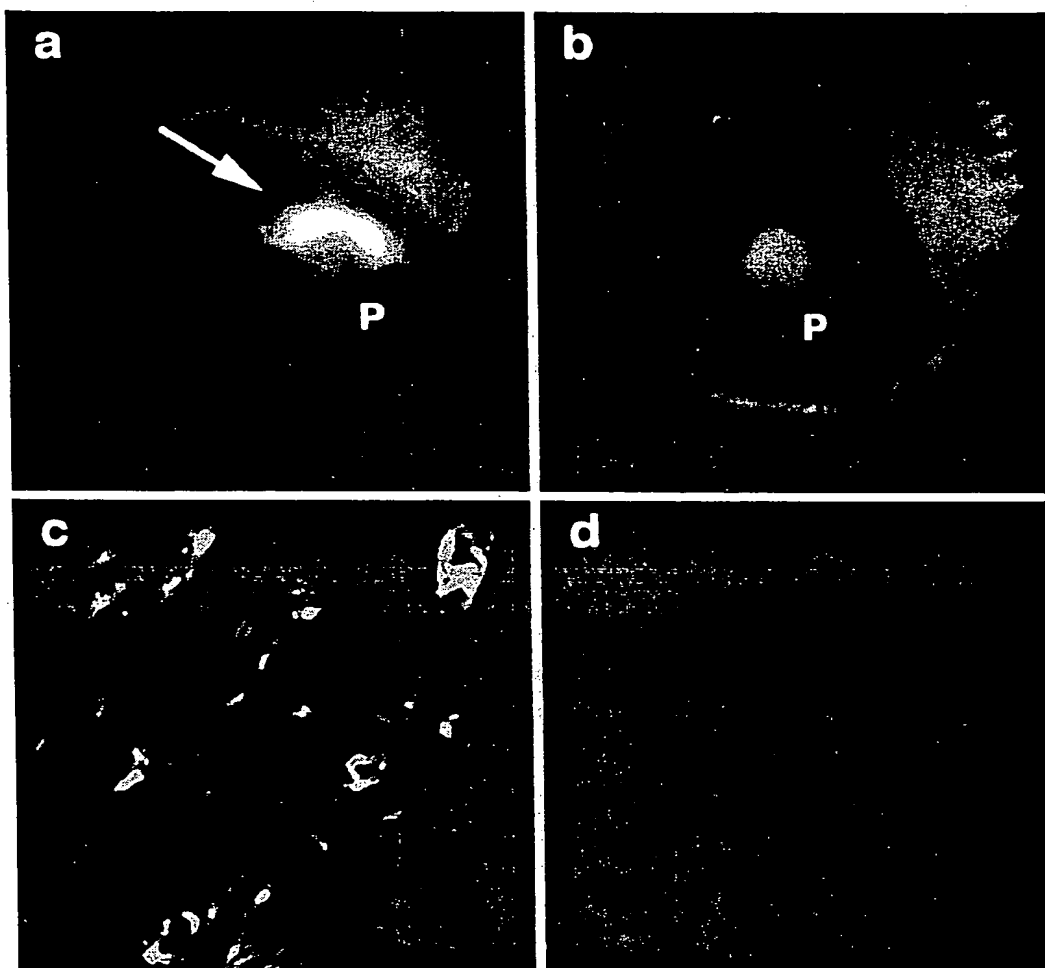


FIG. 10

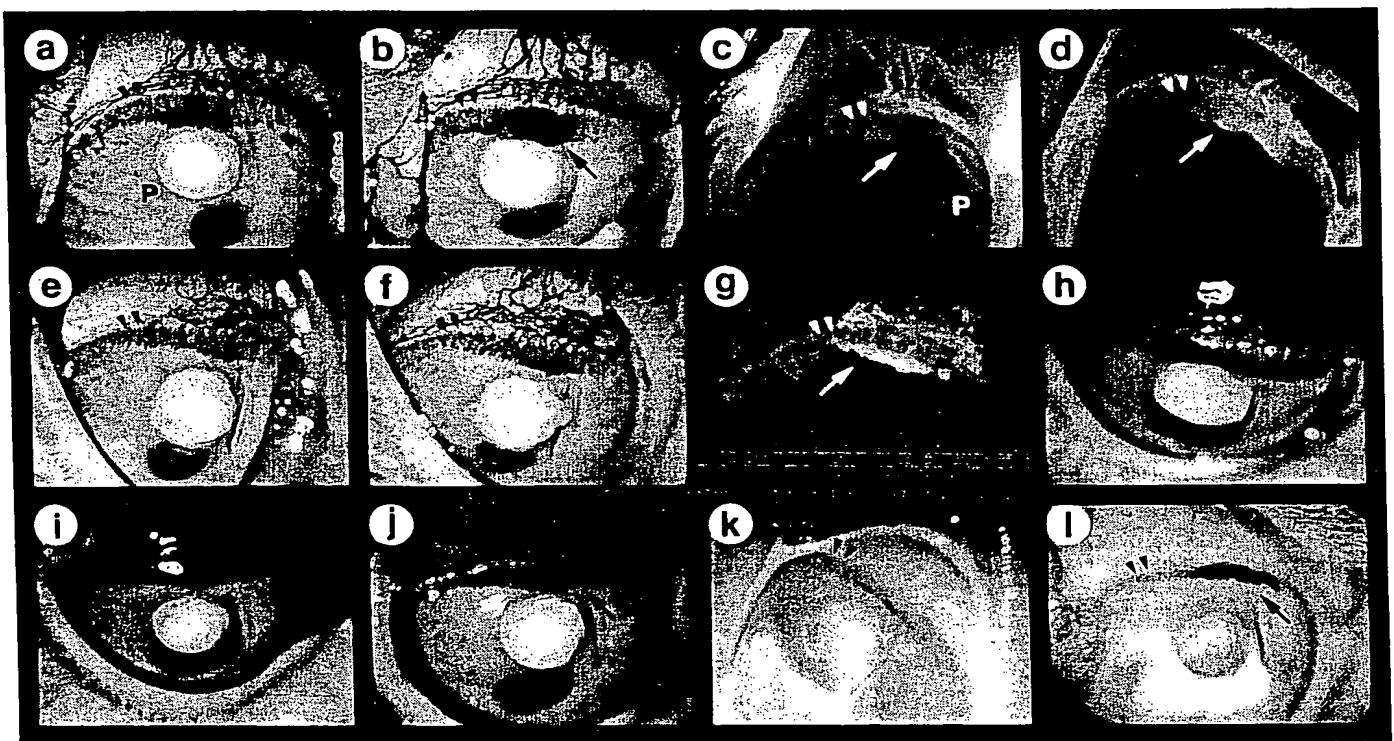


FIG. 11

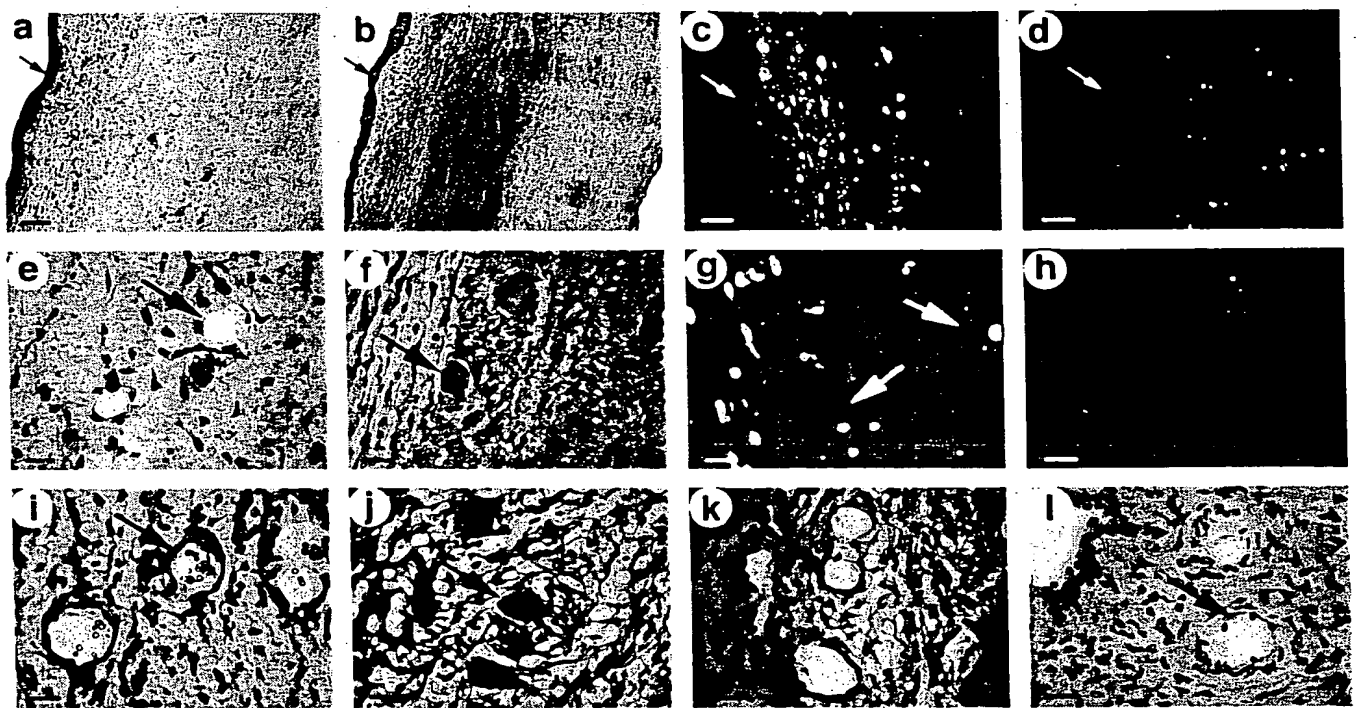


FIG. 12

FIGURE 13

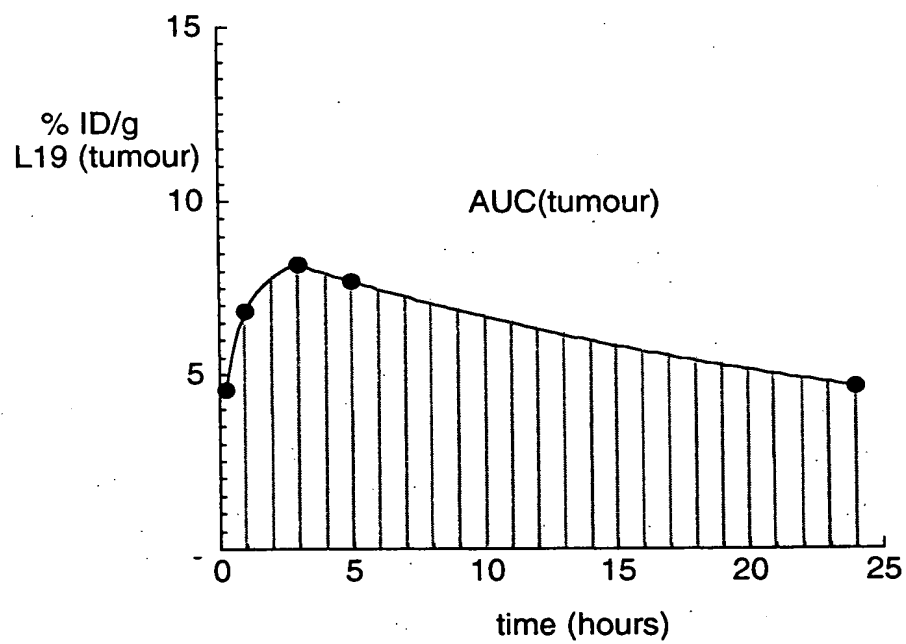
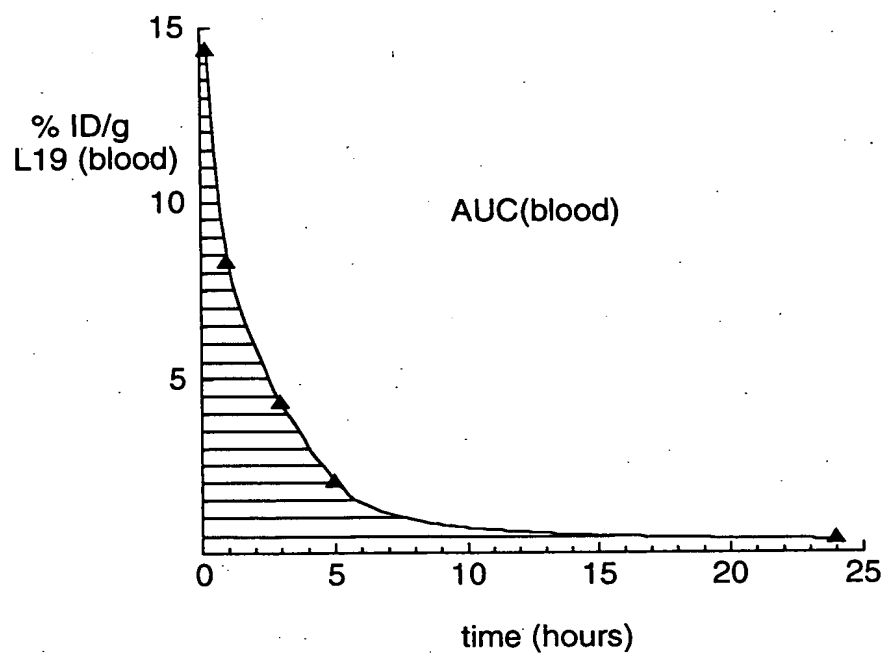


FIGURE 14

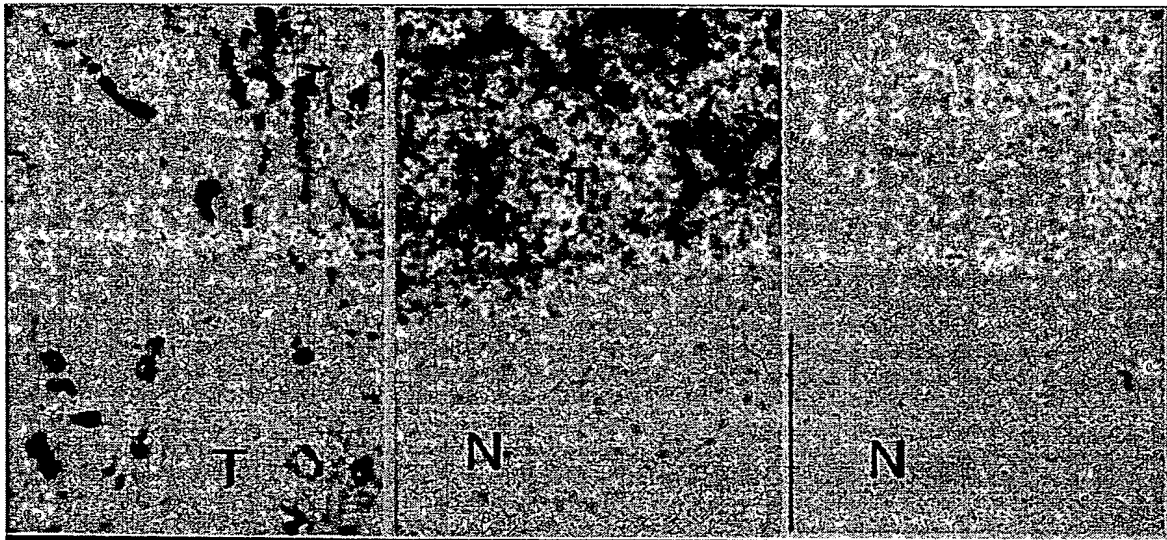
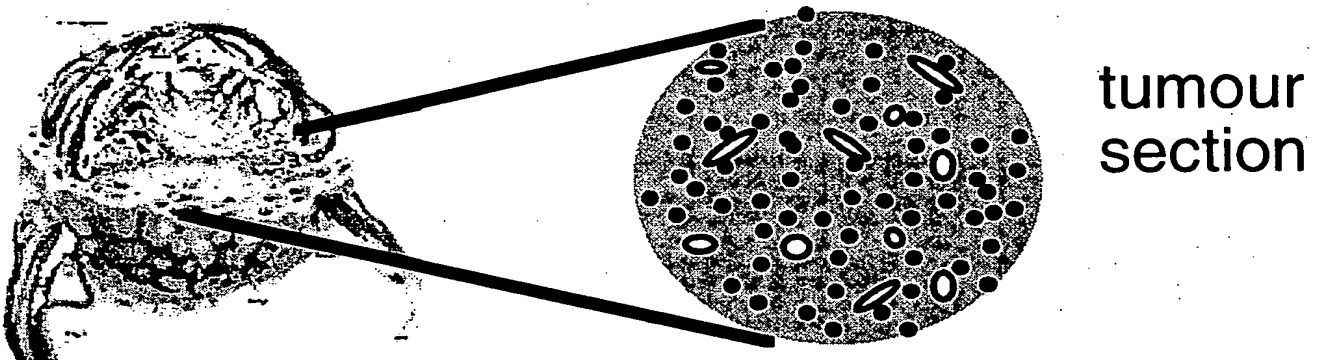


FIGURE 15

Radioimmunotherapy with anti-angiogenesis antibodies



vessels = 0.5 - 5%
of total tumour mass

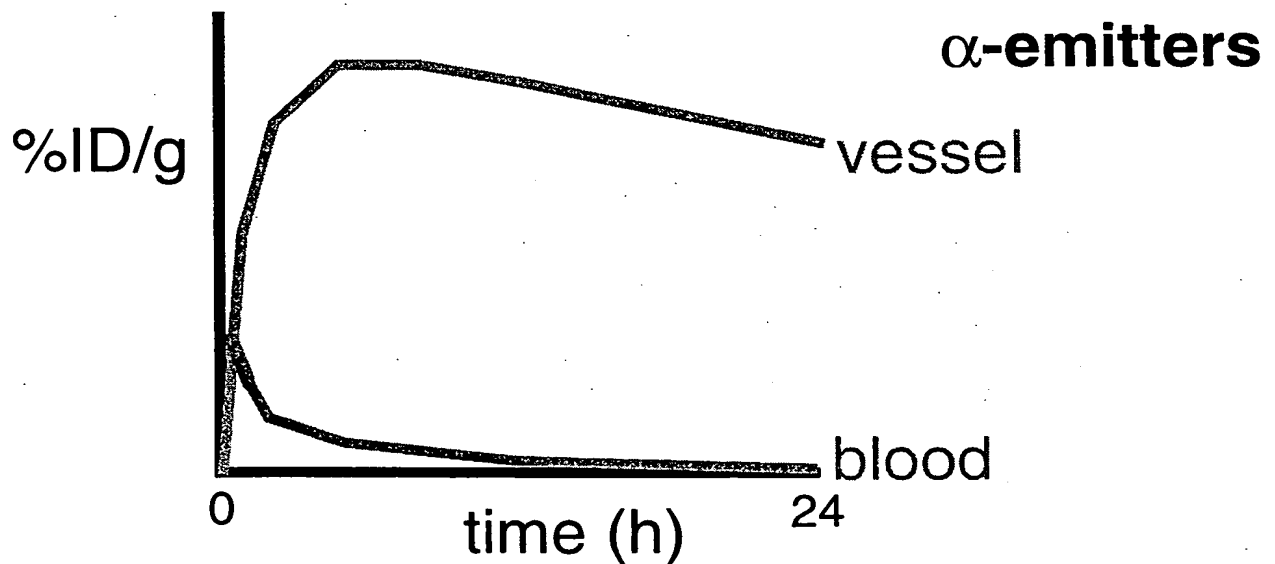
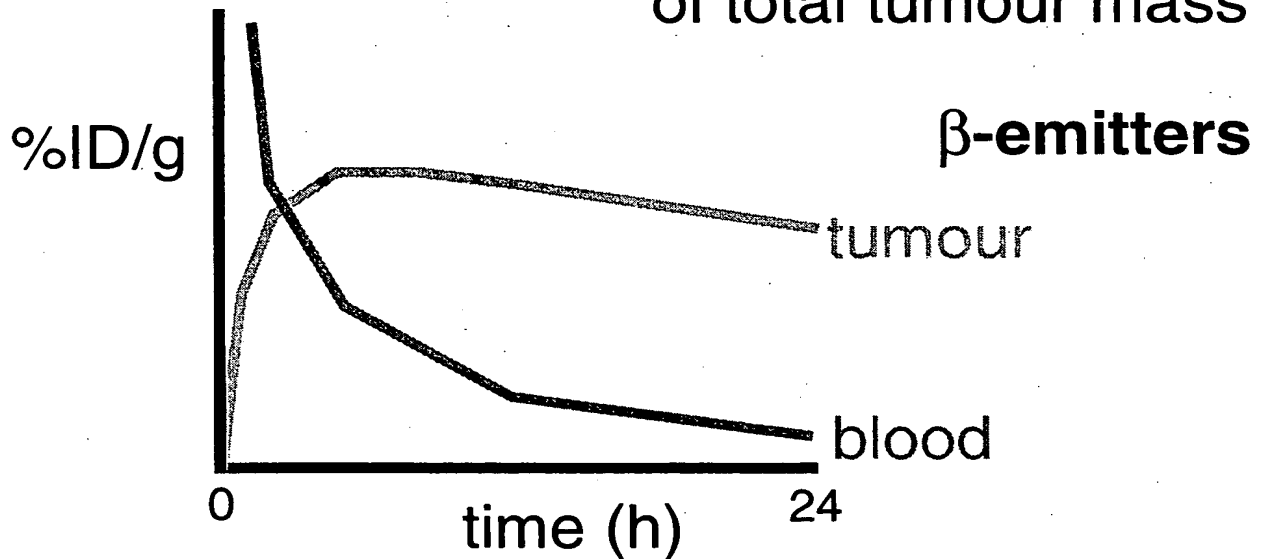


FIGURE 16

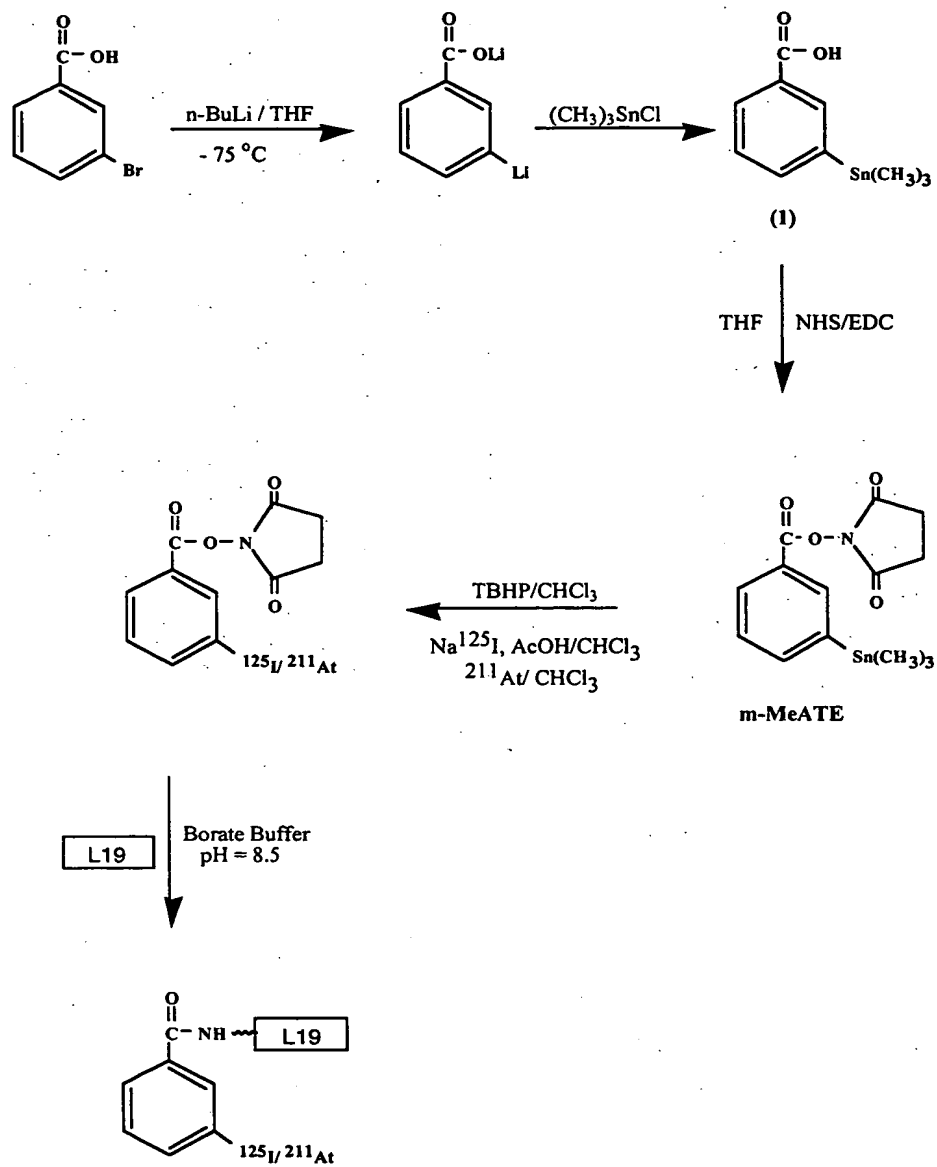
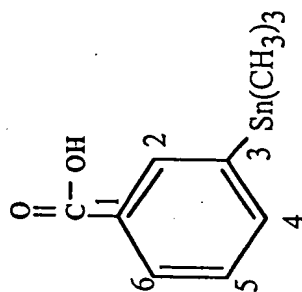


FIGURE 17a

^1H -NMR spectrum of 3-(trimethylstannyl)-benzoic acid in CDCl_3



$(\text{CH}_3)_3$

```

Current Data Parameters
NAME      test_0004
EXPNO     1
PROCNO    1

F2 - Acquisition Parameters
Date_     990804
Time      11.19
INSTRUM   drx600
PROBHD    5 mm TXI 13C
PULPROG   zgpg. eth
TO        8192
SOLVENT   CDCl3
NS        16
DS         2
SWH        9615.385 Hz
FIDRES     1.173753 Hz
AQ         0.4260340 sec
RG         800
DH         52.000 usec
DE         8.00 usec
TE         294.0 K
D1         1.00000000 sec

***** CHANNEL f1 *****
NUC1       1H
P1         8.00 usec
PL1        0.00 dB
PL9        120.00 dB
SFO1       600.1328114 MHz

***** GRADIENT CHANNEL *****
GPMAN1     sine.32
GPX1       0.00 x
GPY1       0.00 x
GPZ1       50.00 x
PZ1        1000.00 usec

F2 - Processing parameters
SI          4096
SF          600.1300239 MHz
WDW         USINE
SSB         3
LB          0.00 Hz
GB          0
PC          1.00

1D NMR plot parameters
CX          20.00 cm
F1P         8.811 ppm
F1          5287.97 Hz
F2P        -0.341 ppm
F2         -204.88 Hz
PPMCH      0.45764 ppm/cm
  
```

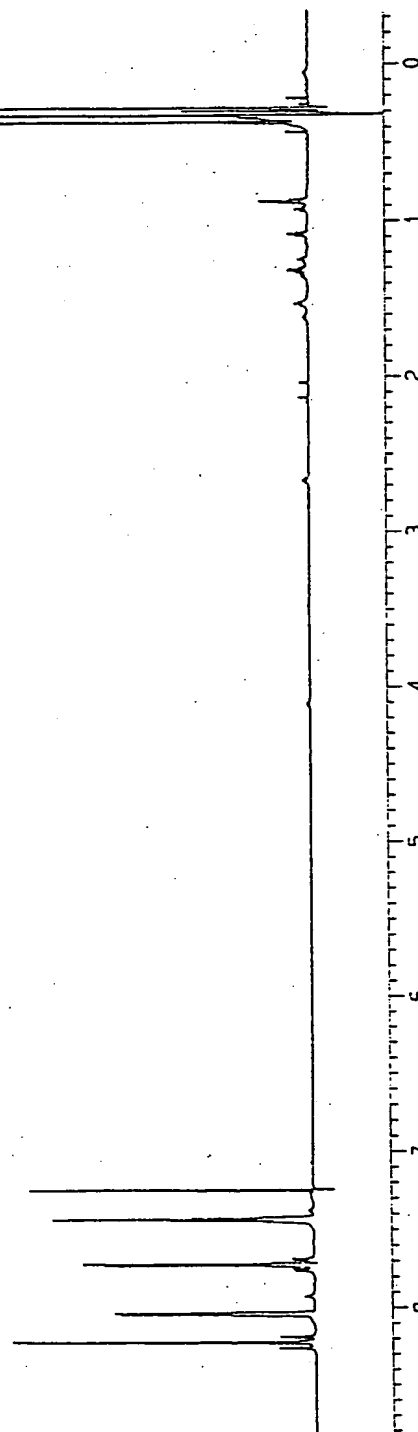
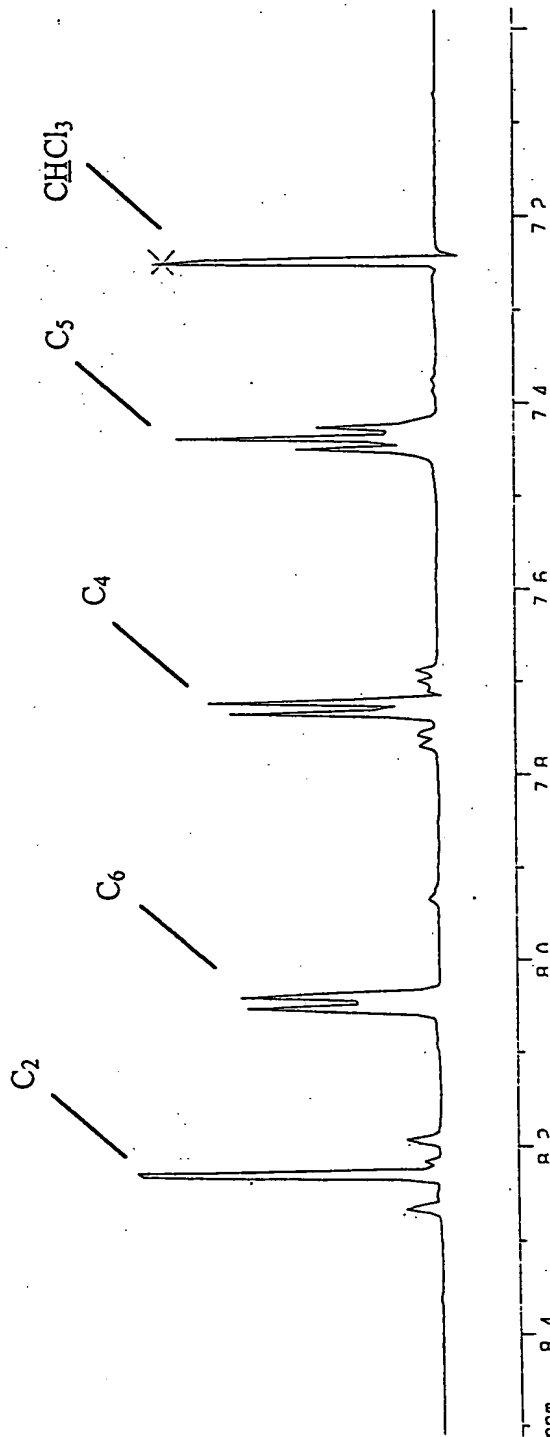
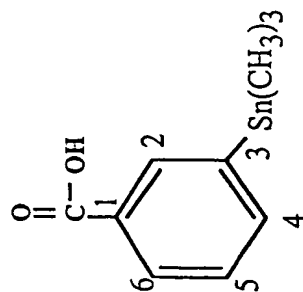


FIGURE 17b

^1H -NMR spectrum of 3-(trimethylstannyl)-benzoic acid in CDCl_3



Current Data Parameters
NAME test_0004
EXPNO 1
PROCNO 1

F2 - Acquisition Parameters
Date_ 990804
Time 11.19
INSTRUM drx600
PROBHD 5 mm TXI 13C
PULPROG zgpg. eth
TD 8192
SOLVENT CDCl3
NS 16
DS 2
SWH 9615.385 Hz
FIDRES 1.173753 Hz
AQ 0.4260340 sec
RG 800
DM 52.000 usec
DE 8.00 usec
TE 294.0 K
D1 1.00000000 sec

----- CHANNEL f1 -----
NUC1 ^1H
P1 8.00 usec
PL1 0.00 dB
PL9 120.00 dB
SFO1 600.1326114 MHz

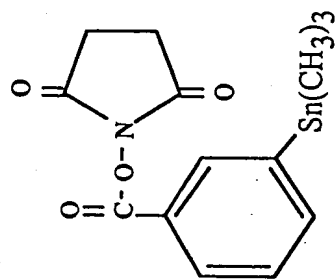
----- GRADIENT CHANNEL -----
GPMAX1 sine.32
GPX1 0.00 %
GPY1 0.00 %
GPZ1 50.00 %
PZ1 1000.00 usec

F2 - Processing parameters
SI 4096
SF 600.1300239 MHz
WDW OSINE
SSB 3
LB 0.00 Hz
GB 0
PC 1.00

1D NMR plot parameters
CX 20.00 cm
FIP 8.513 ppm
F1 5108.78 Hz
F2 6.977 ppm
F2 4187.17 Hz
PPHCH 0.07678 ppm/cm

FIGURE 18a

^1H -NMR spectrum of N-succinimidyl-3-(trimethylstannyl)-benzoate in CDCl_3



m-MeATE

$(\text{CH}_3)_3$

$\text{CO}-\text{CH}_2-\text{CH}_2-\text{CO}$

C_6
 C_4
 C_5
 CHCl_3

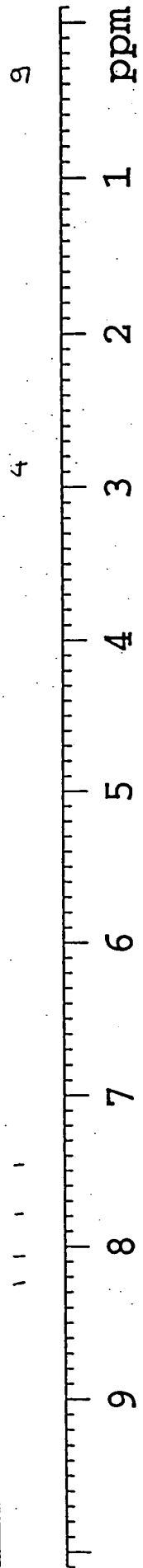


FIGURE 18b

^1H -NMR spectrum of N-succinimidy-3-(trimethylstannyl)-benzoate in CDCl_3

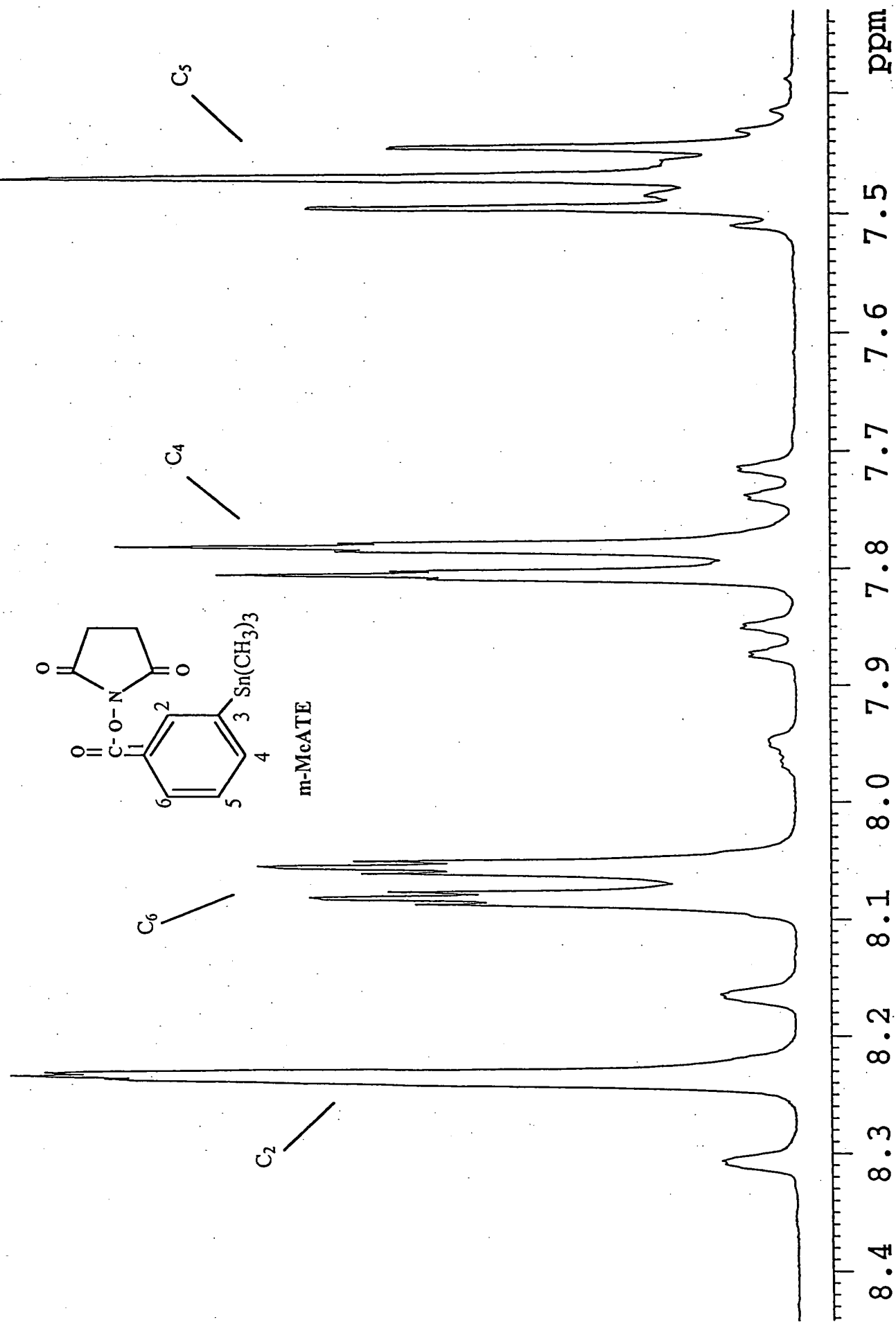


FIGURE 19

EI-MS of N-succinimidyl-3-(trimethylstannyl)-benzoate in CDCl₃

